

FIG. 1(a)

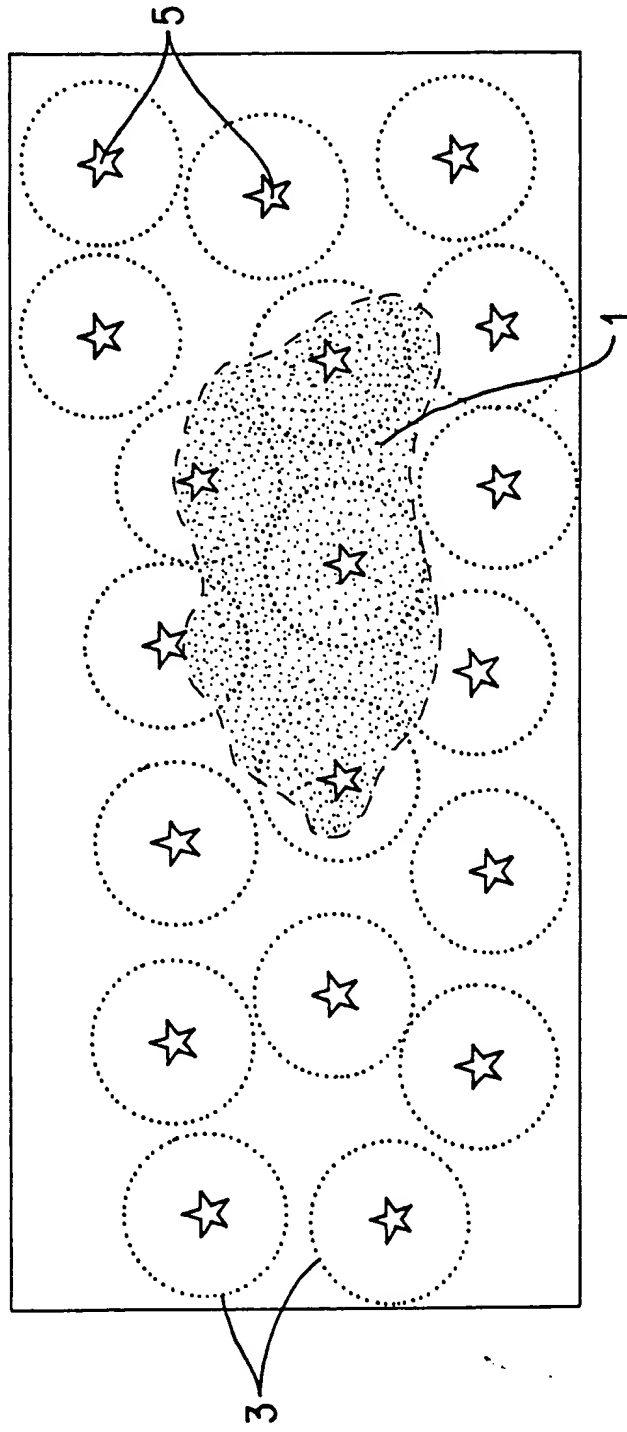


FIG. 1(b)

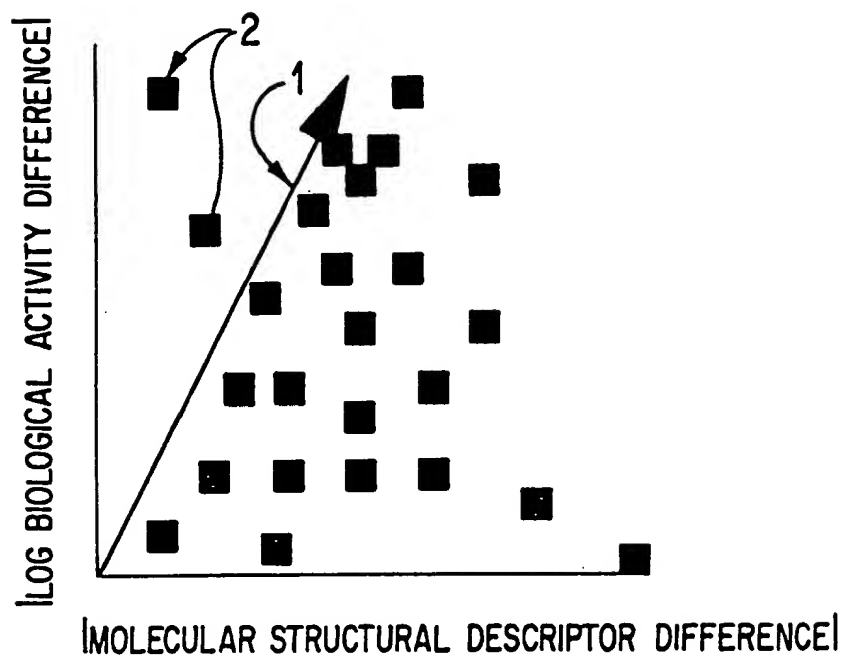


FIG. 2

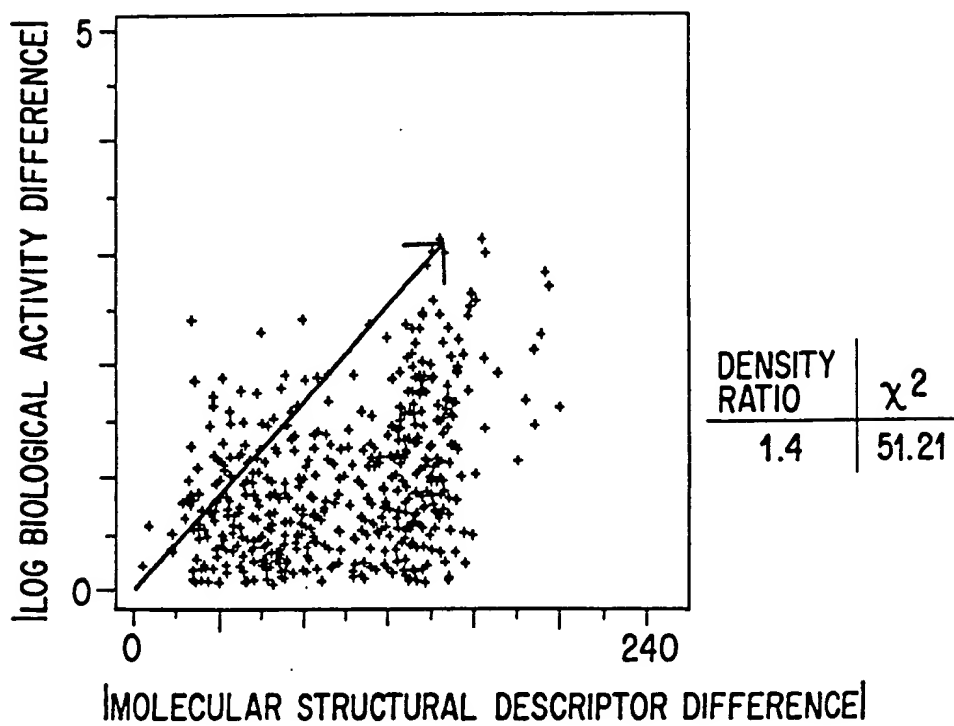


FIG. 3

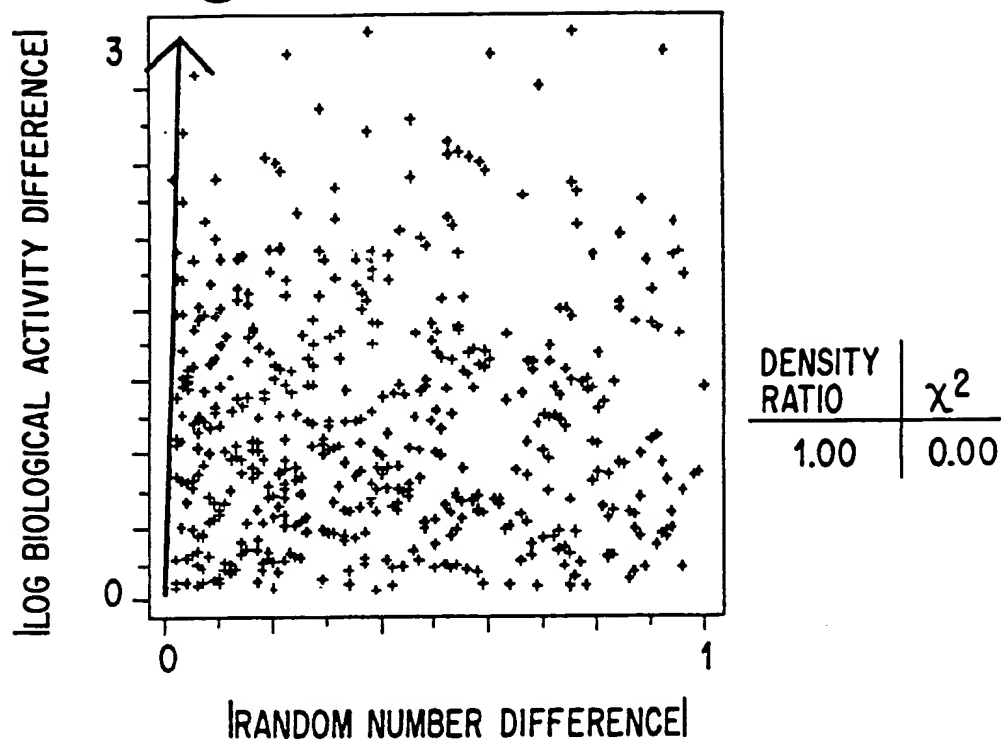


FIG. 4

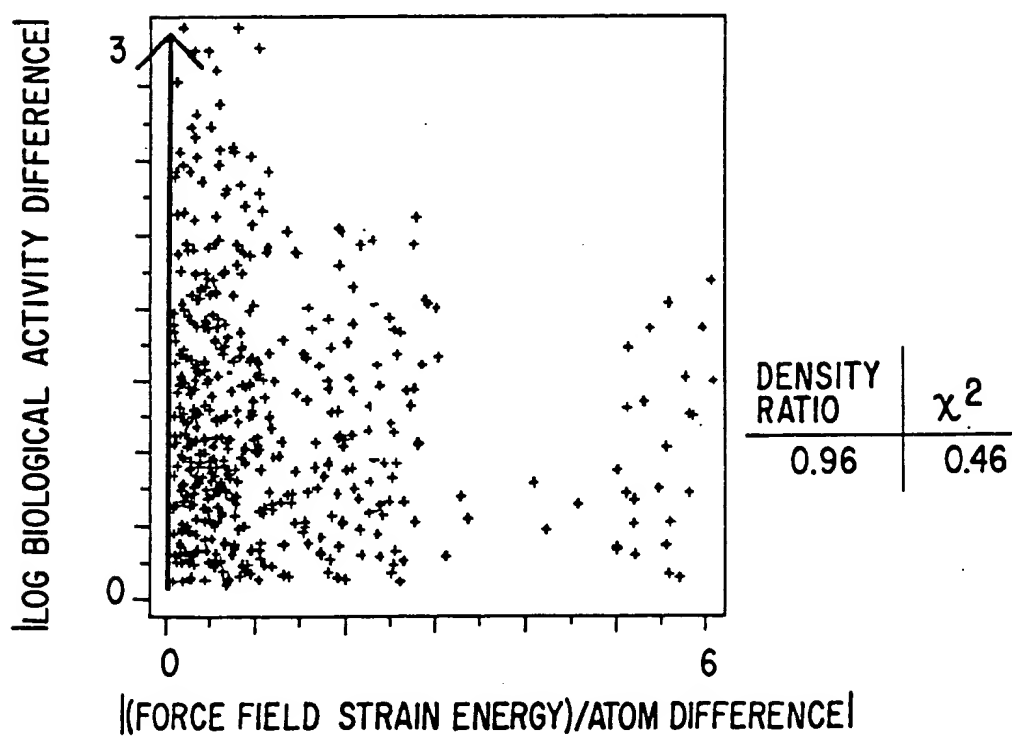
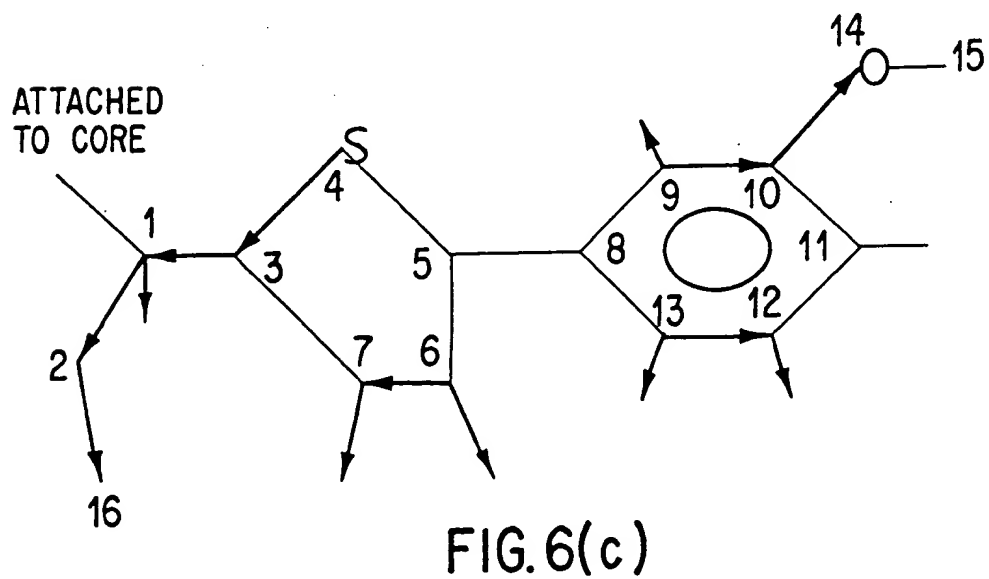
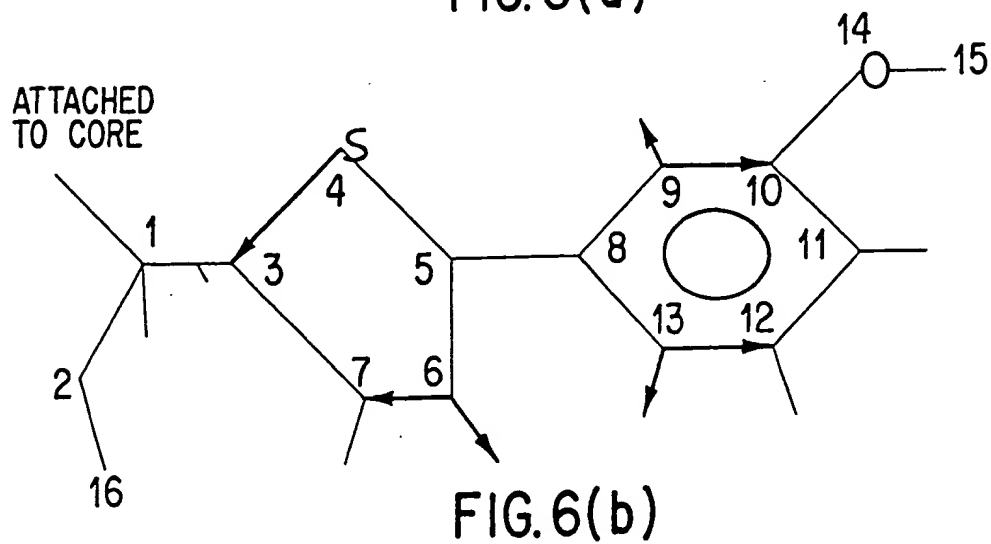
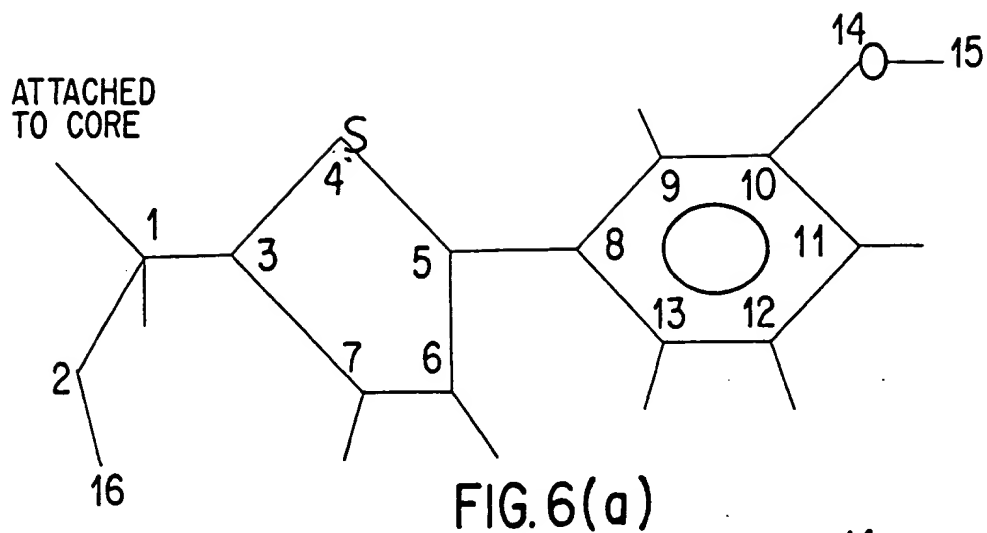


FIG. 5



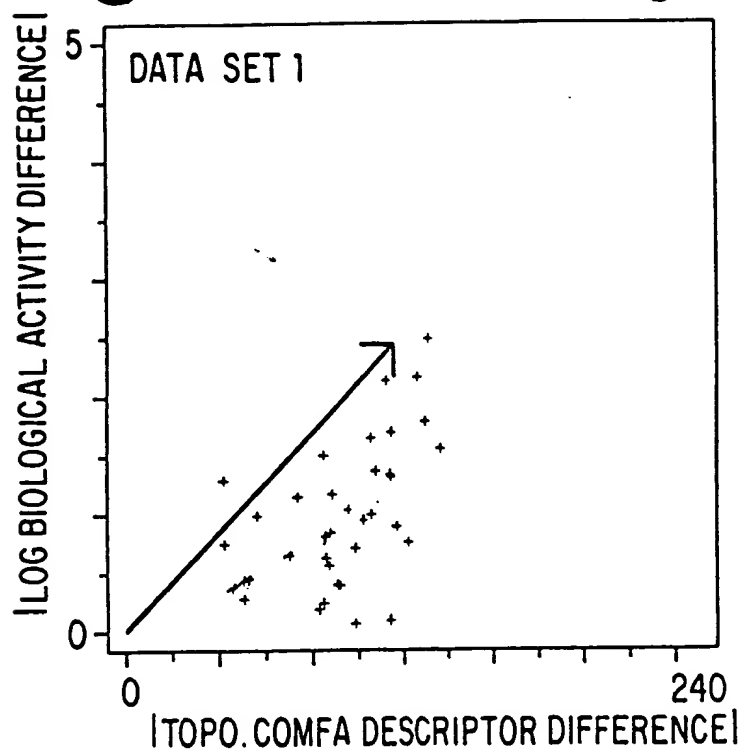


FIG.7(a)

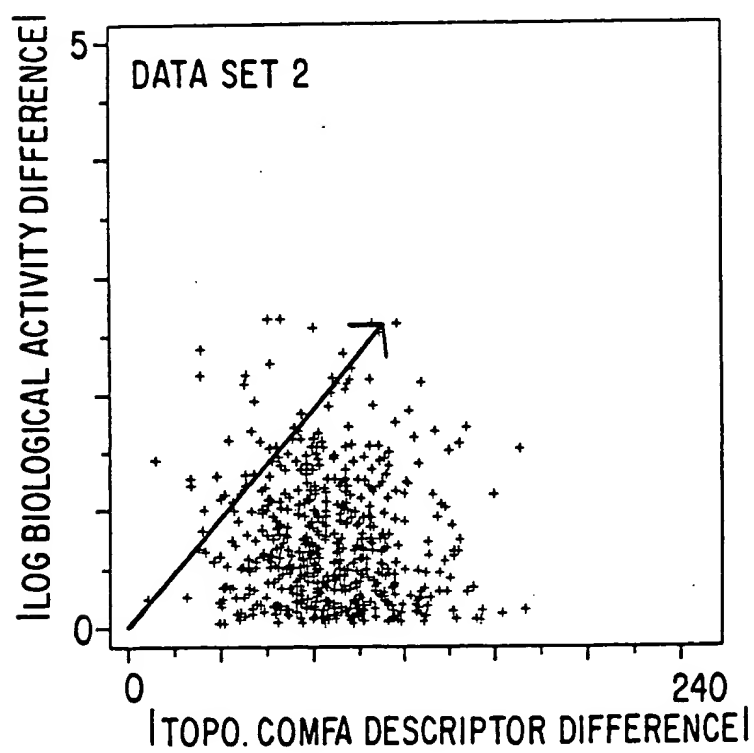


FIG. 7(b)

FIG. 7(d)

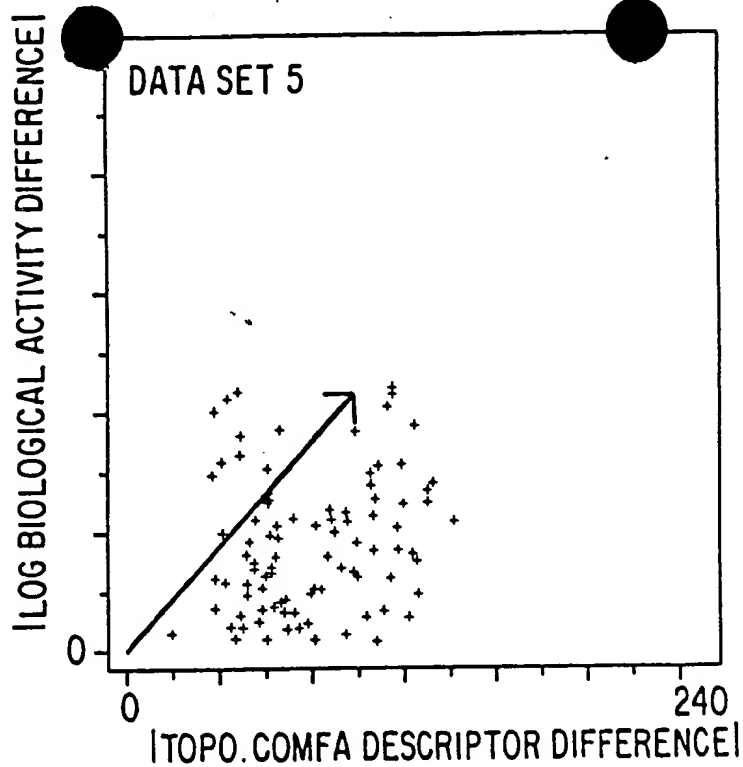


FIG.7(e)

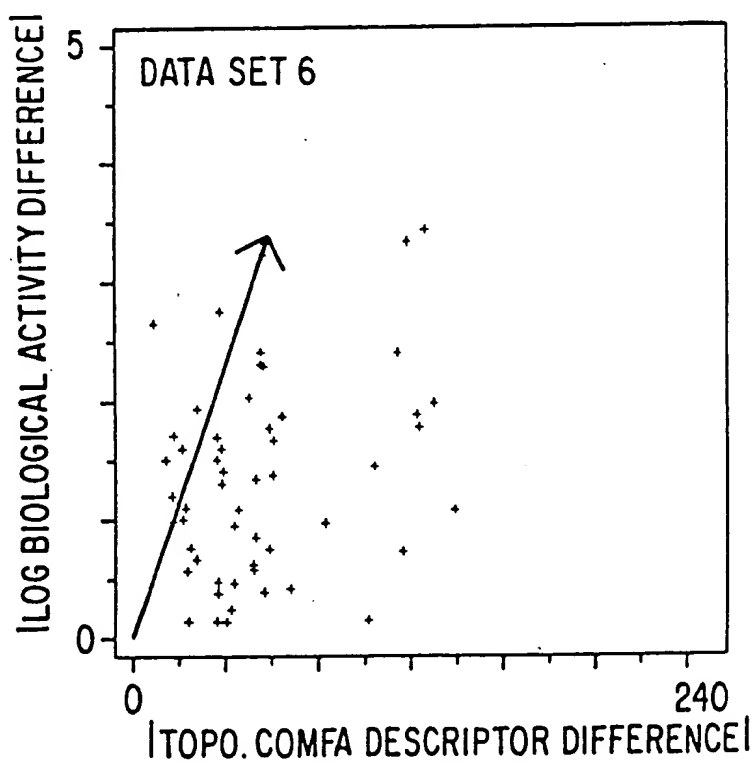


FIG.7(f)

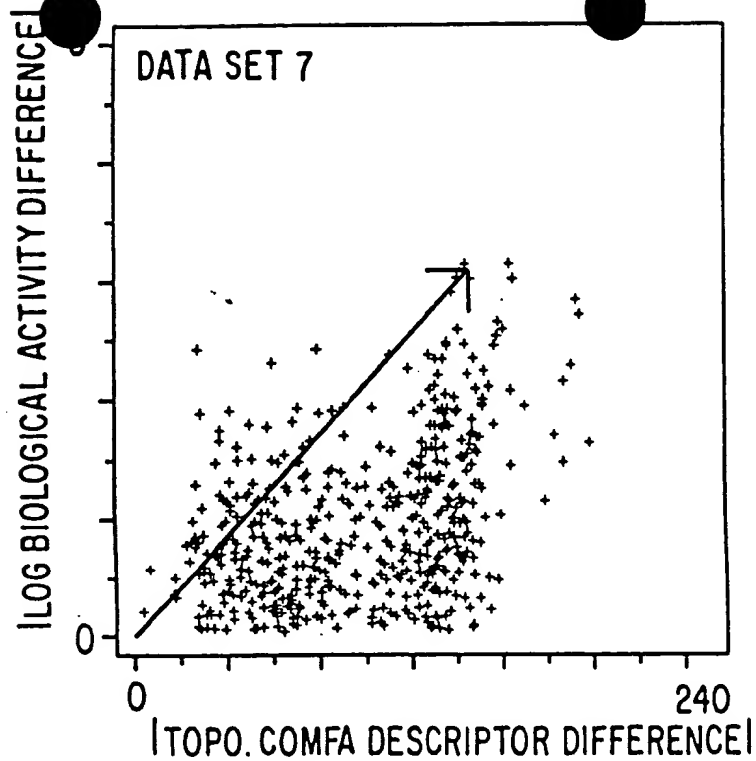


FIG.7(g)

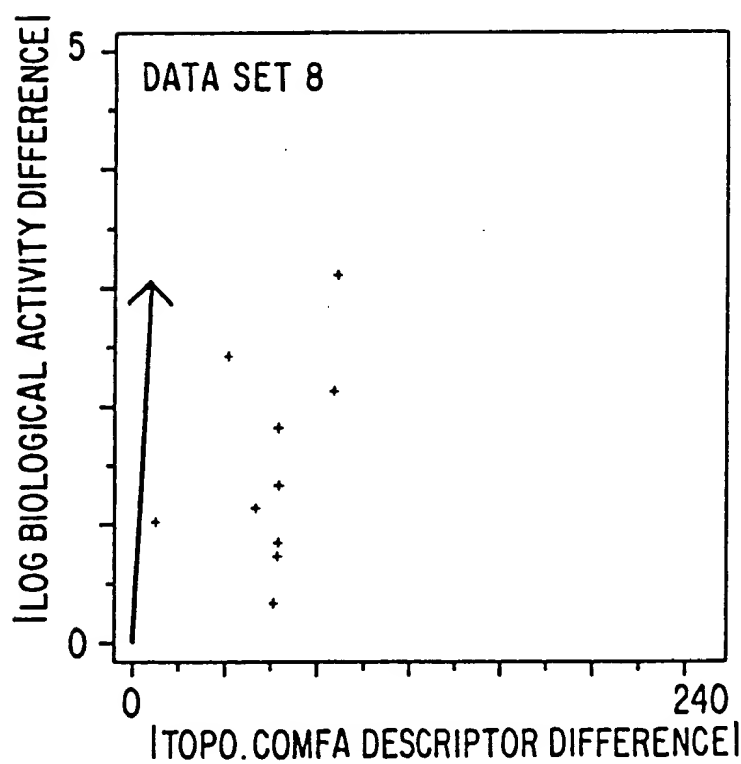


FIG.7(h)

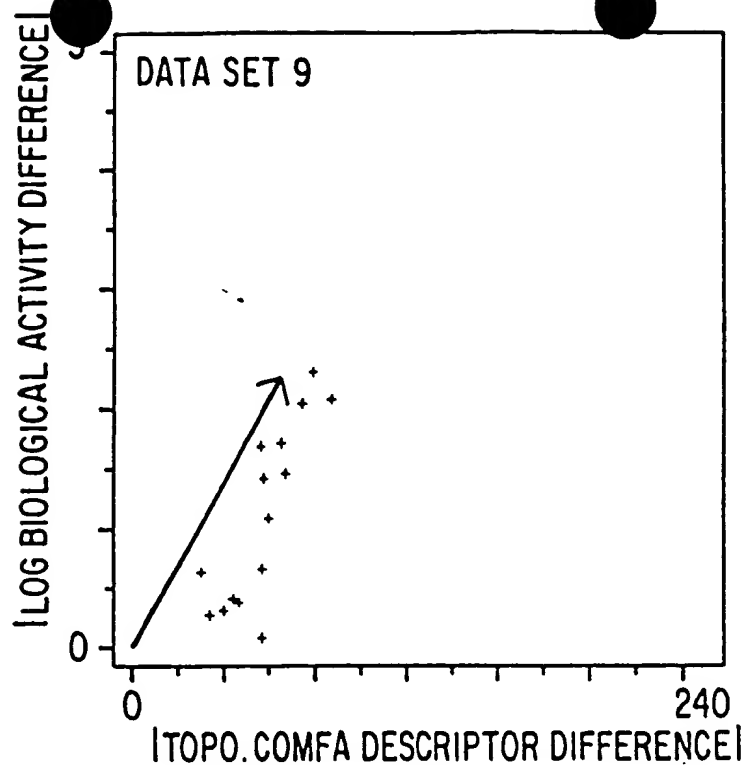


FIG.7(i)

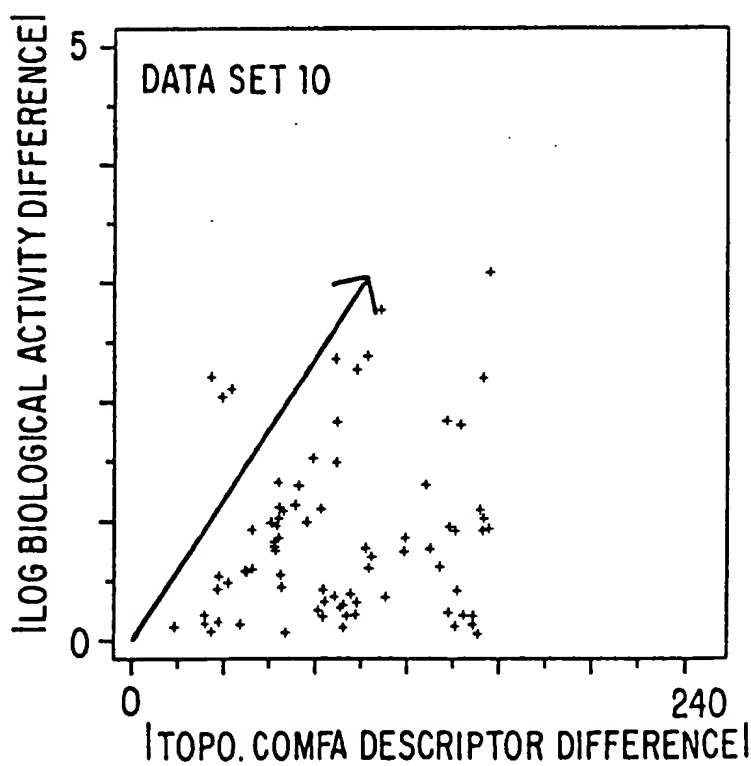


FIG.7(j)

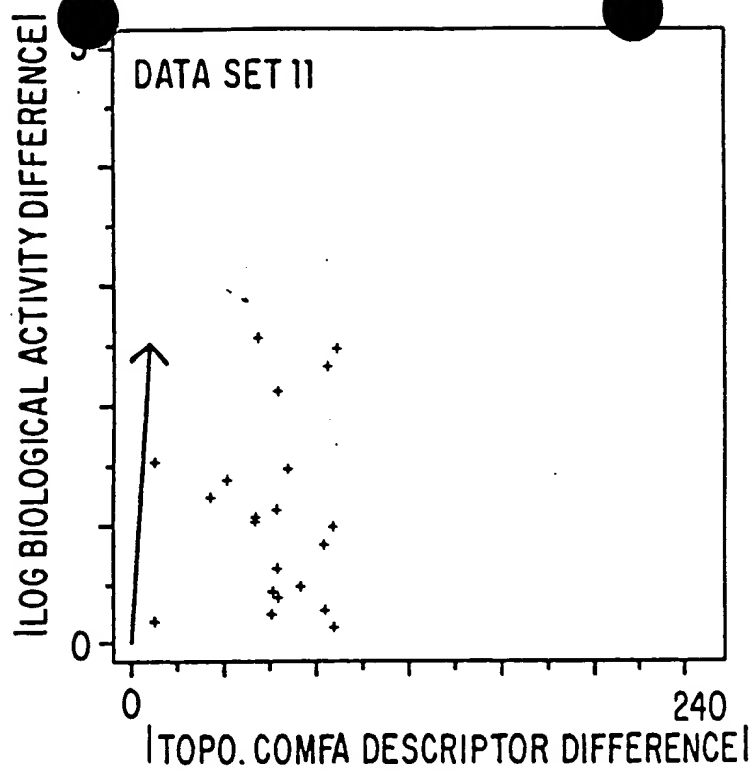


FIG.7(k)

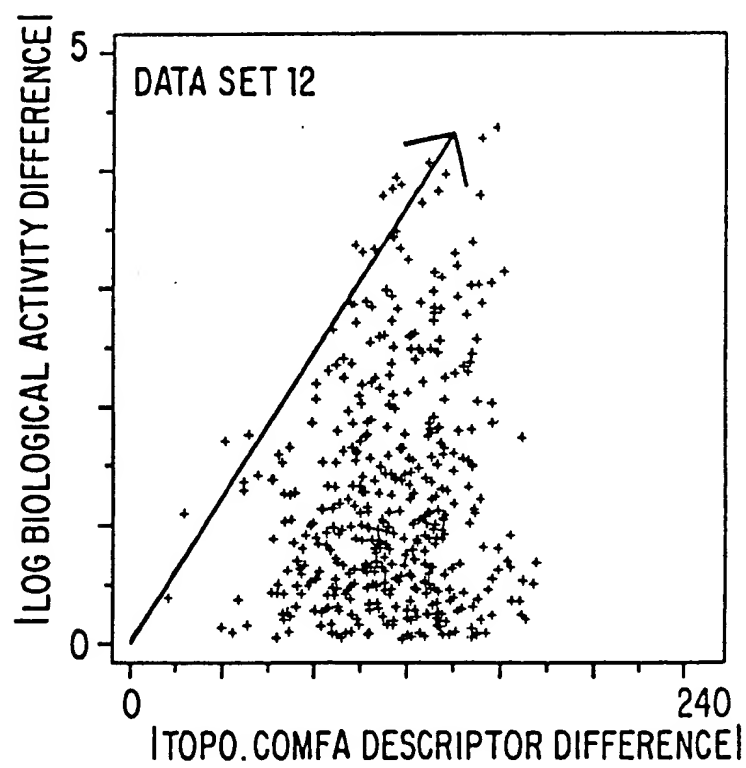


FIG. 7(1)

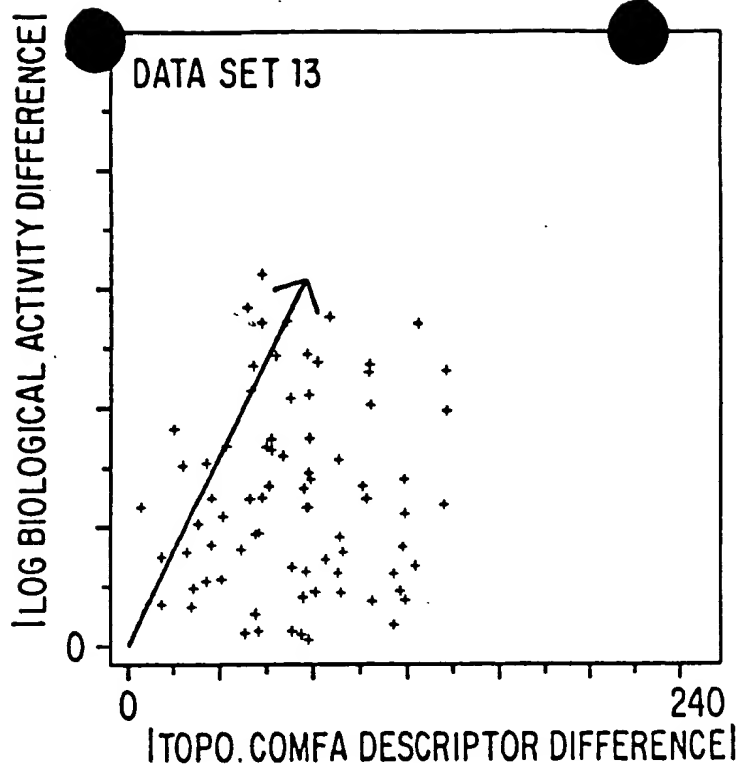


FIG.7(m)

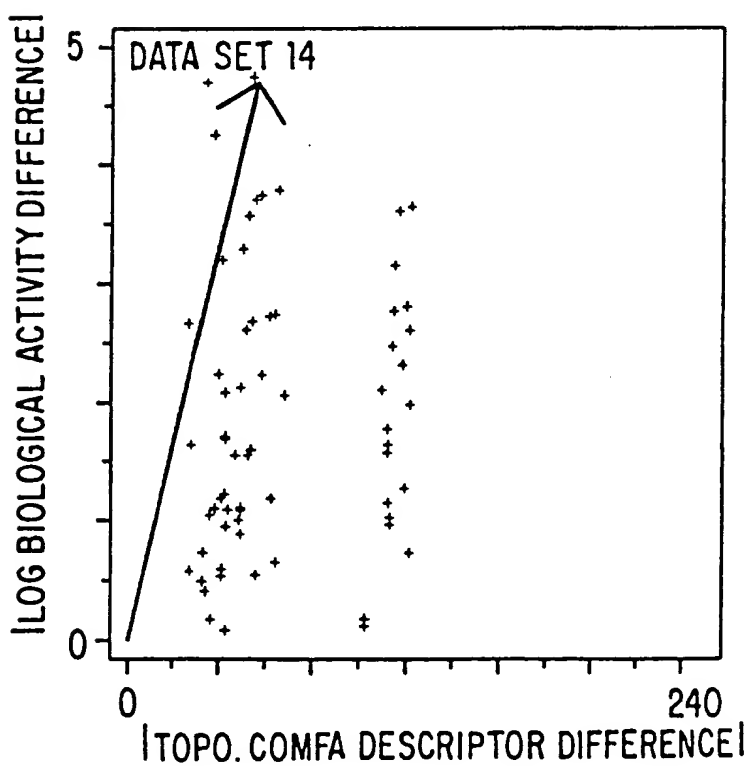


FIG.7(n)

DATA SET 15

Y-axis: |LOG BIOLOGICAL ACTIVITY DIFFERENCE|

X-axis: |TOPO.COMFA DESCRIPTOR DIFFERENCE|

0 240

FIG. 7(p)

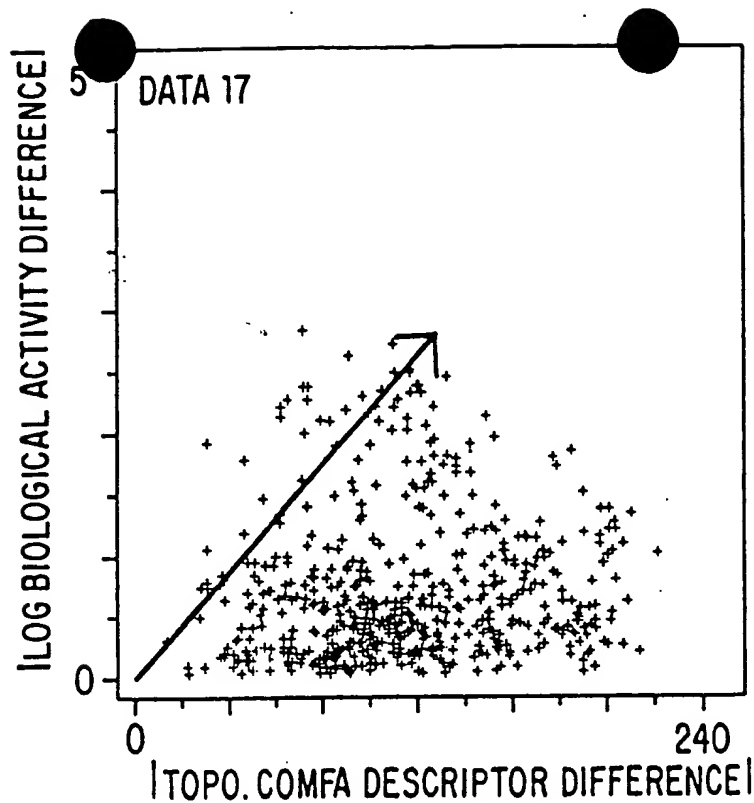


FIG.7(q)

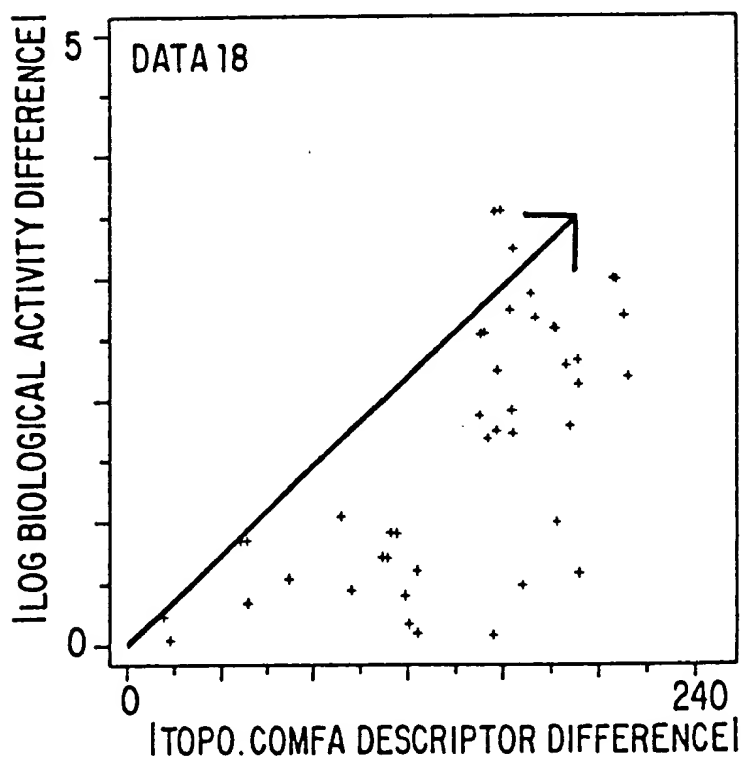


FIG.7(r)

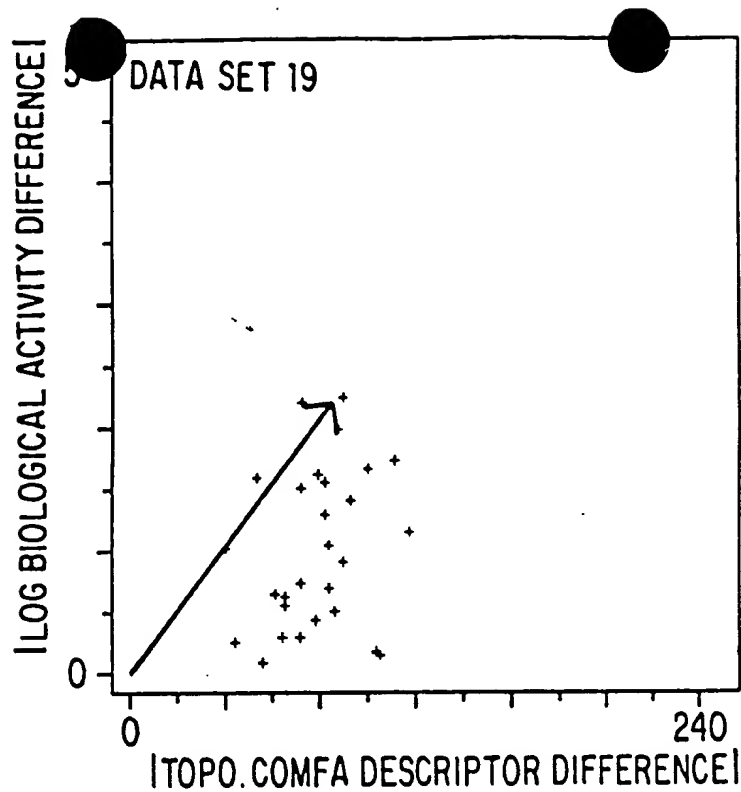


FIG.7(s)

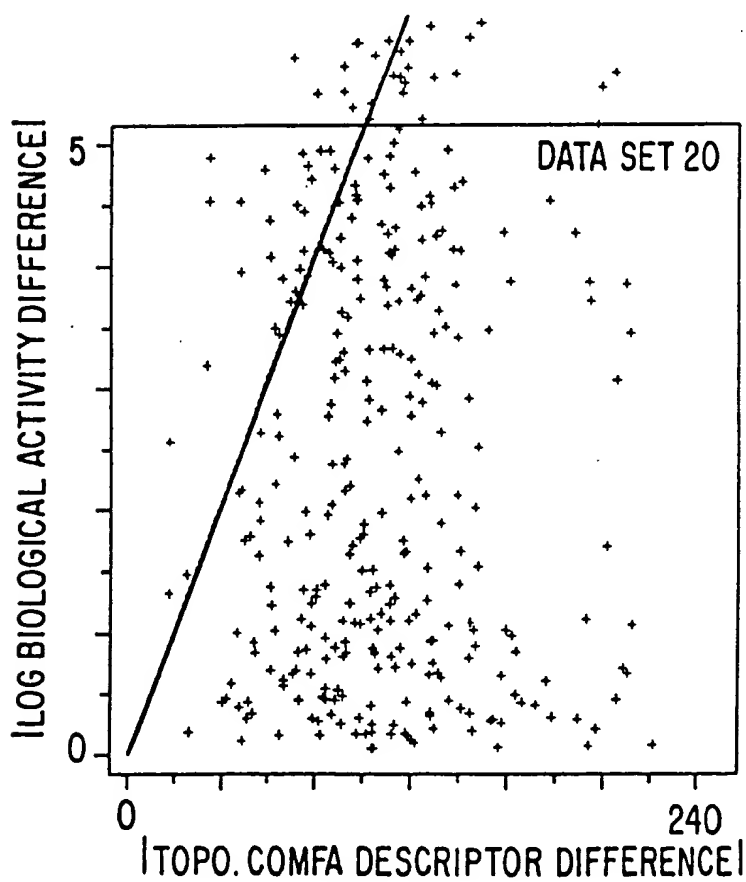
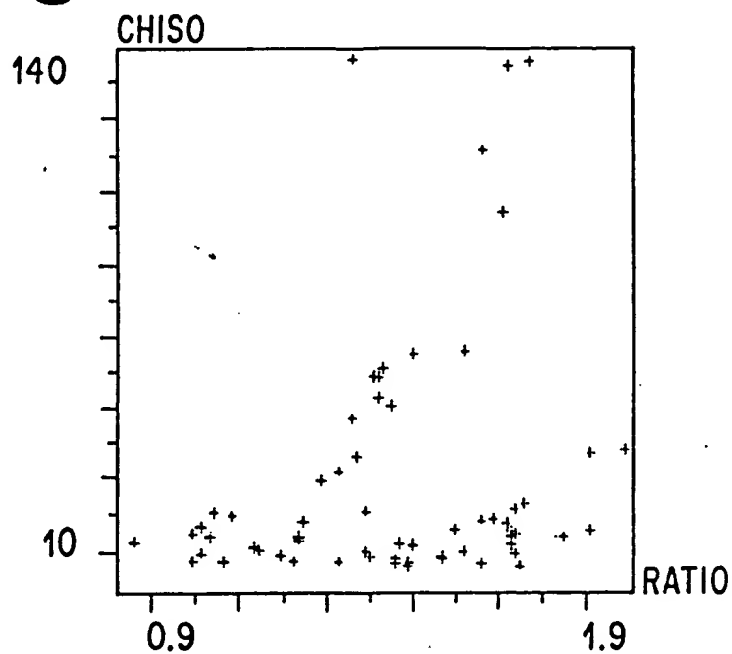
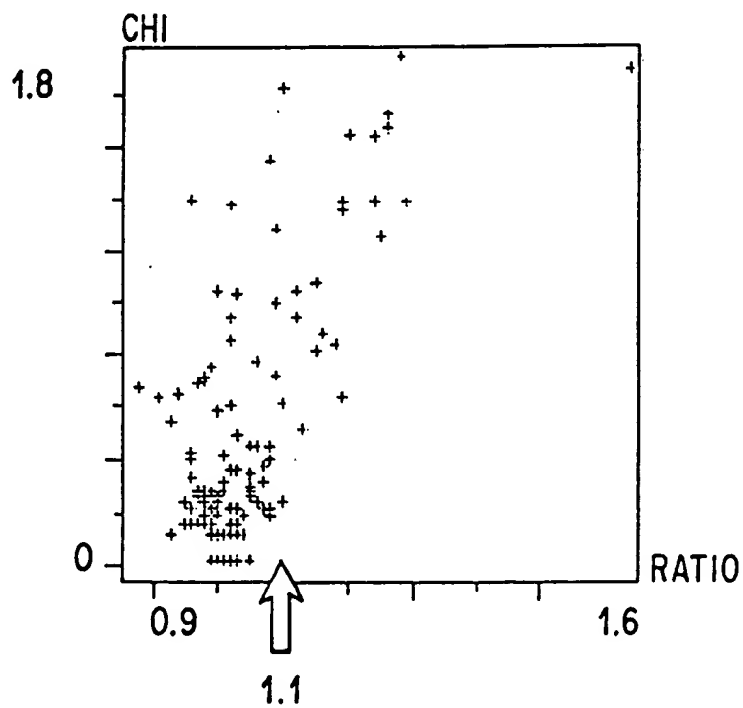


FIG.7(t)



1.1

FIG. 8(a)



1.1

FIG. 8(b)

FOUOED" F F 29260

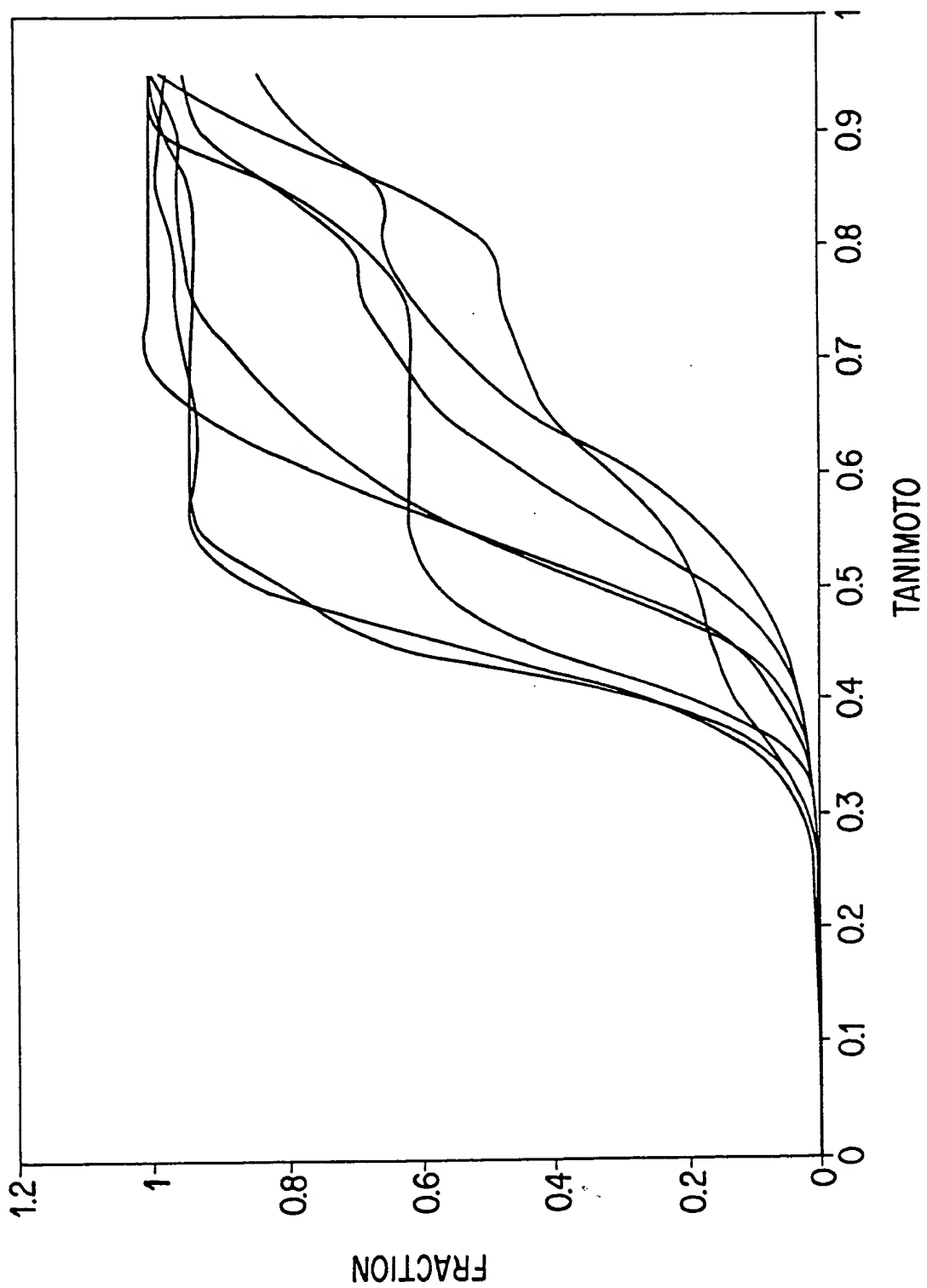


FIG. 9(a)

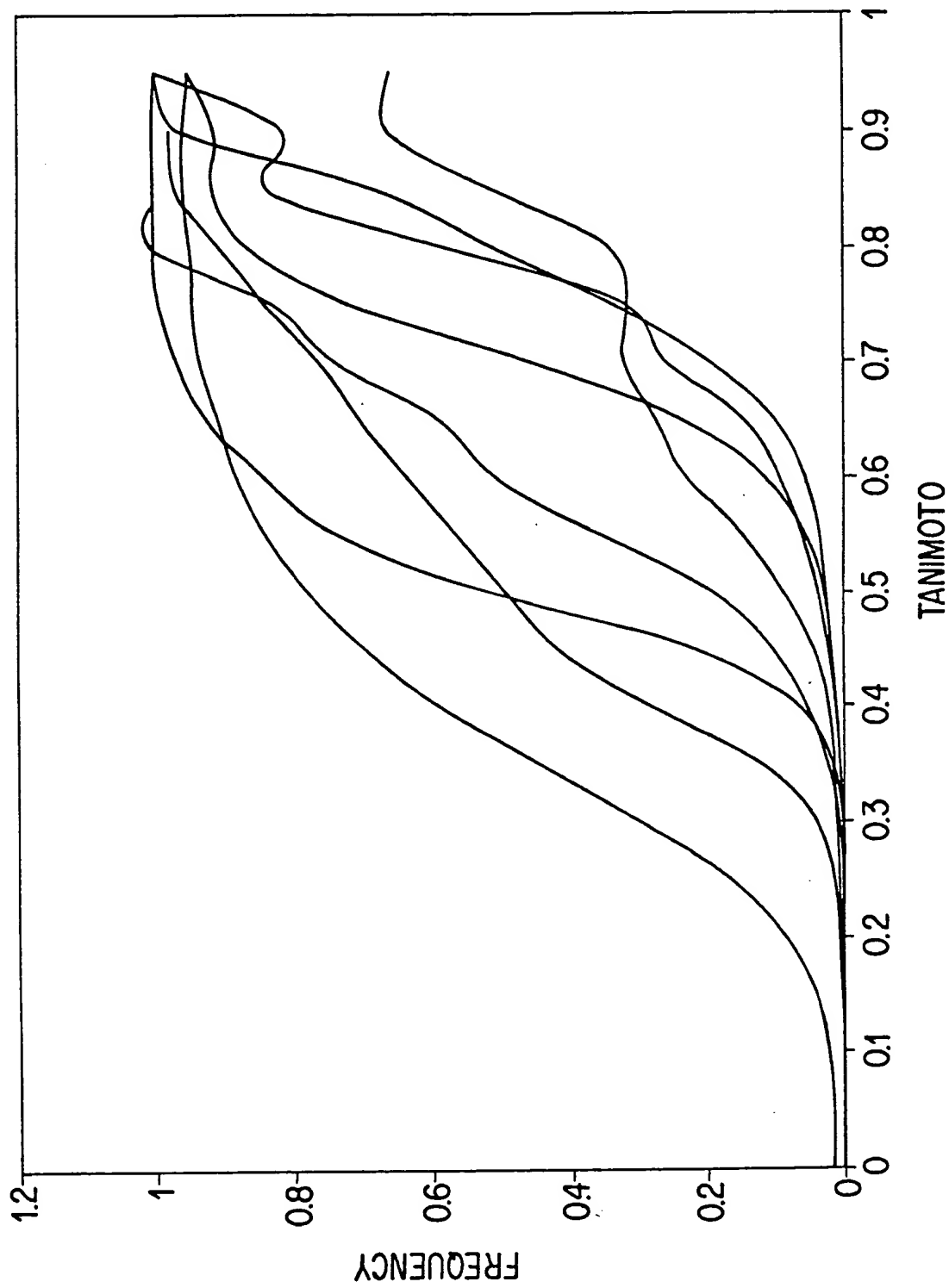
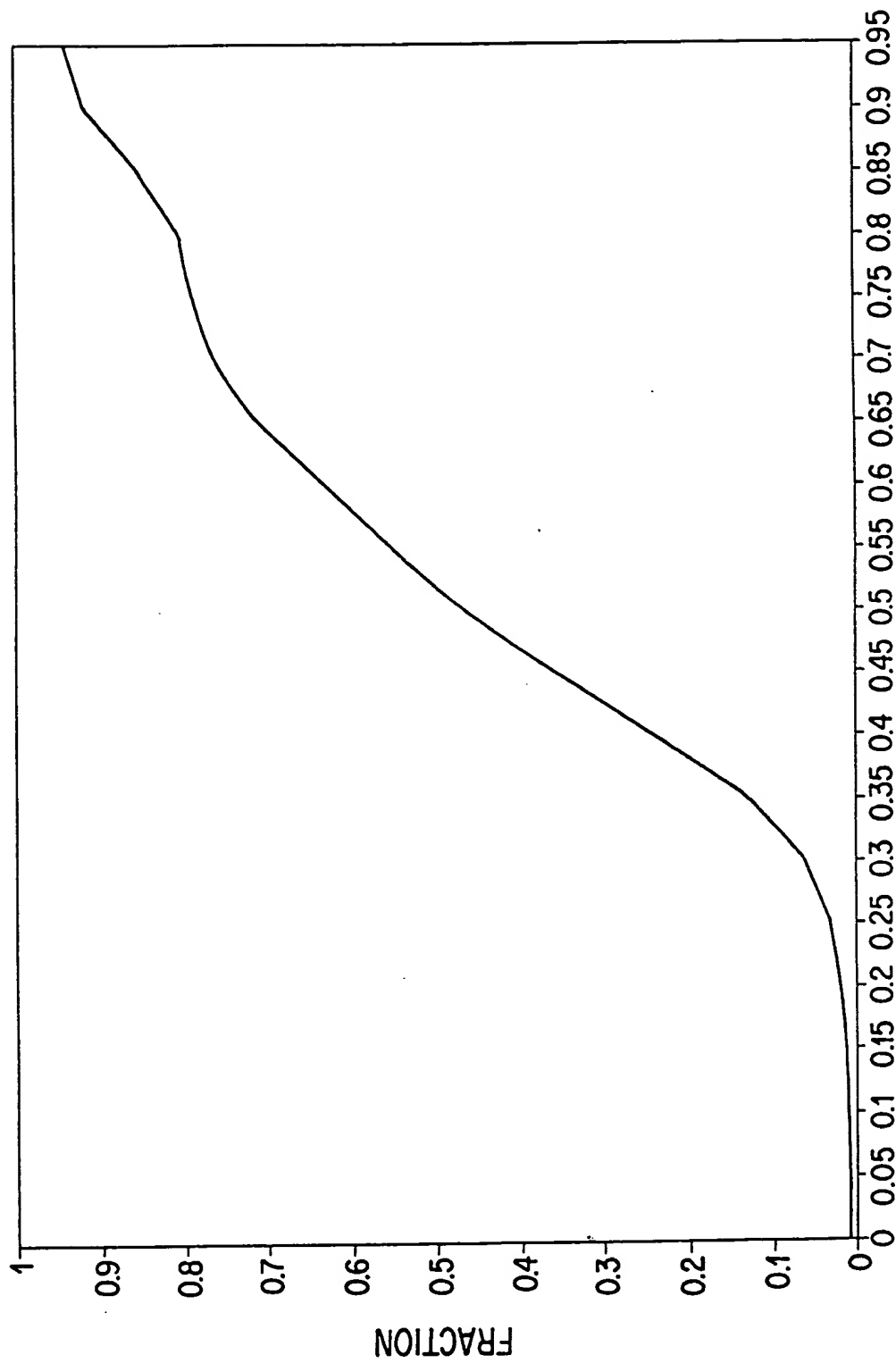


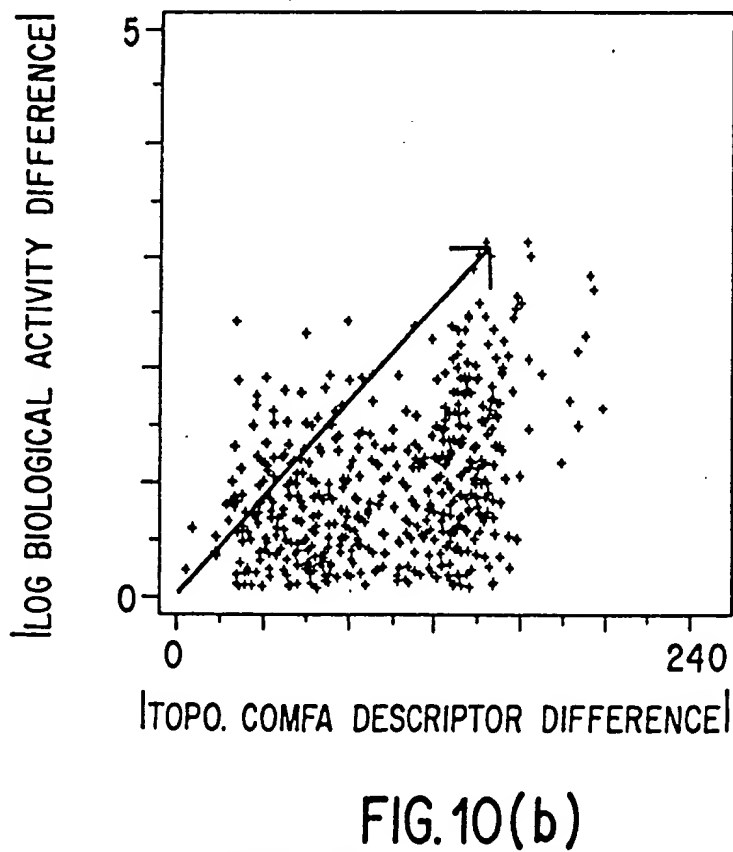
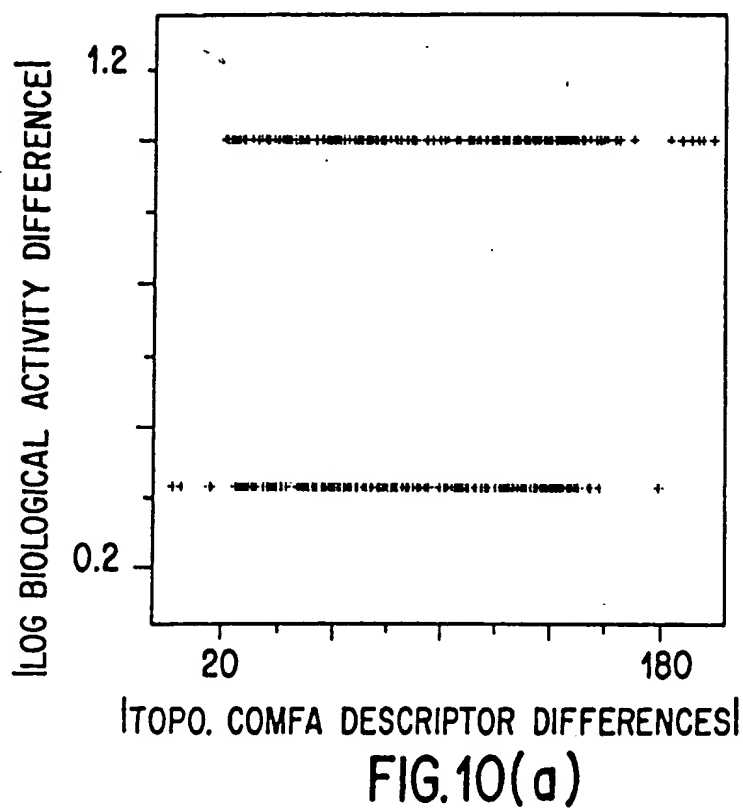
FIG. 9(b)

T09020" T29460



TANIMOTO

FIG.9(c)



00541-0004
T05020" T49460

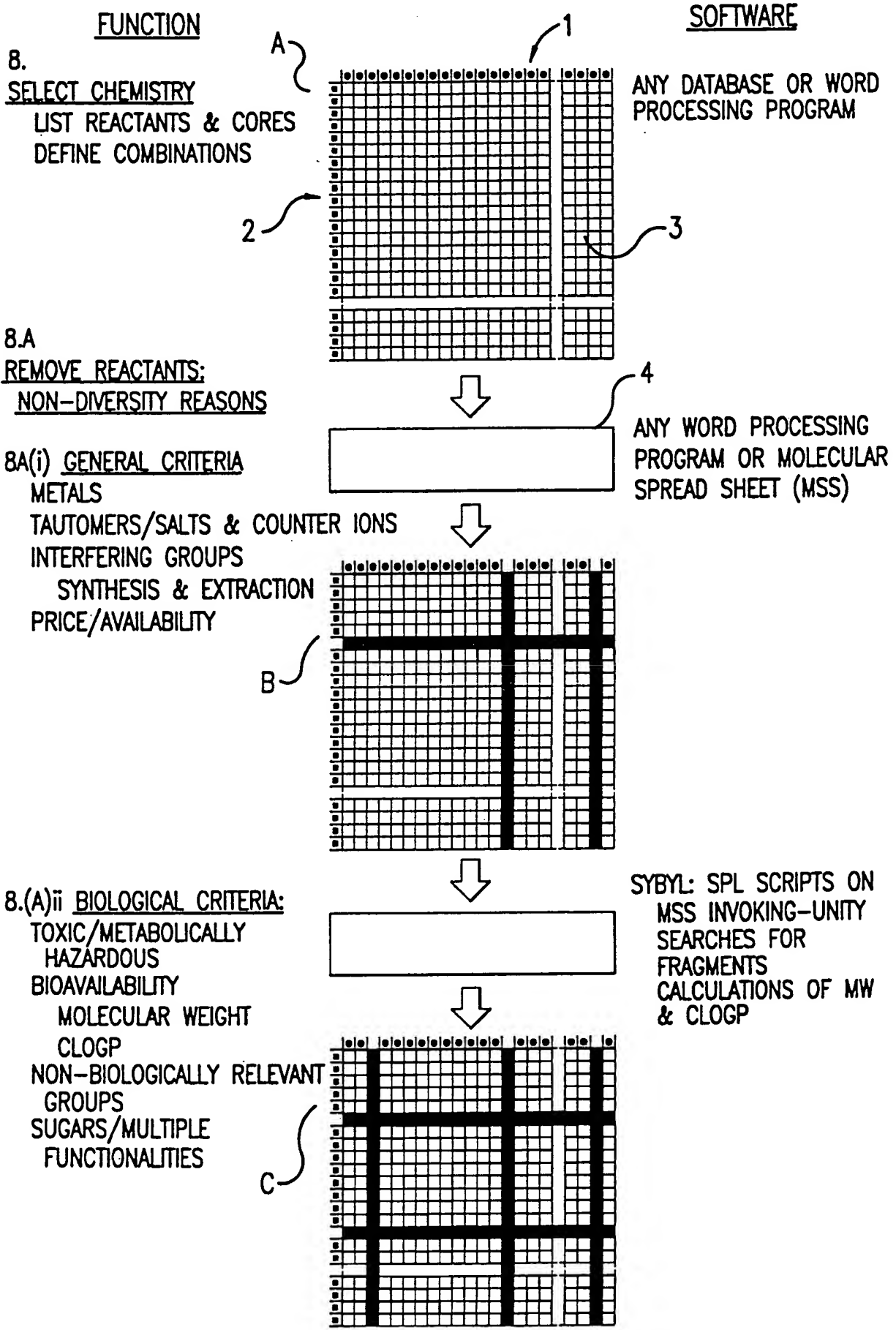


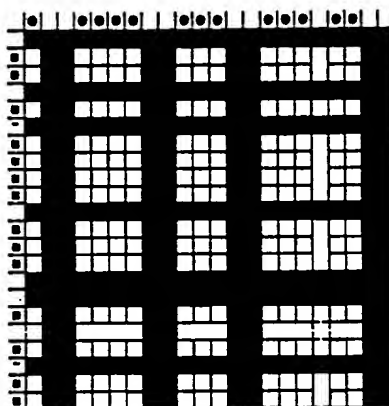
FIG.11(a)

8B

REMOVE NON-DIVERSE REACTANTS:

3D STRUCTURE GENERATION
 TOPOMERIC CONFORMER
 ALIGNMENT
 CoMFA FIELD GENERATION:
 HYDROGEN BOND FIELD
 GENERATION
 ROTATABLE BOND FIELD
 ATTENUATION
 CALCULATE FIELD
 DIFFERENCES
 CLUSTER
 REACTANT SELECTION FROM
 EACH CLUSTER

D



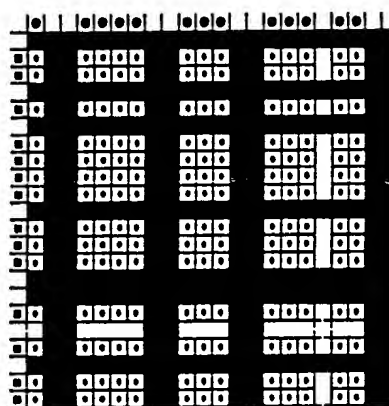
SYBYL: SELECTOR &
 CONCORD 3.2.1
 SPECIALIZED SOFTWARE:
 APPENDIX "A"
 (SYBYL: CoMFA
 STERIC & MSS)

SYBYL: HIERCHICAL
 CLUSTER - MSS
 SYLBL: MSS

8C

COMBINE REACTANTS TO BUILD PRODUCTS

E



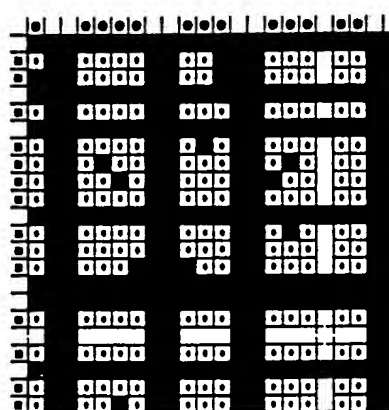
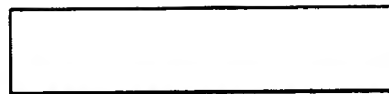
SYBYL: LEGION MSS

8D

REMOVE PRODUCTS:
NON-DIVERSITY REASONS

GENERAL CRITERIA:

F



SYBYL: SPL SCRIPTS
 ON MSS

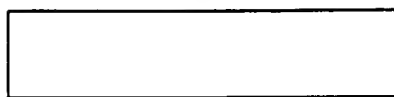


FIG.11(b)

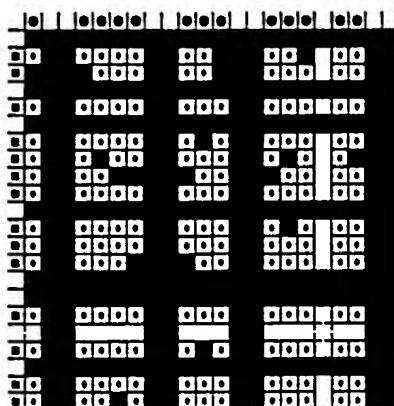
09794 0994
 T05020 T09460

8D

BIOLOGICAL CRITERIA:
METABOLICALLY HAZARDOUS
BIOAVAILABILITY
MOLECULAR WEIGHT
CLOGP



G

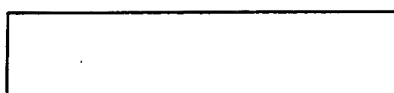


SYBYL: SPL SCRIPTS ON
MSS INVOKING -
UNITY SEARCHES FOR
FRAGMENTS
CALCULATIONS OF MW
& CLOGP

8E

REMOVE NON-DIVERSE
PRODUCTS:

GENERATE TANIMOTO
2D FINGERPRINTS
SAMPLE PRODUCTS:
EXCLUDE TAN ≥ 0.85



UNITY



H

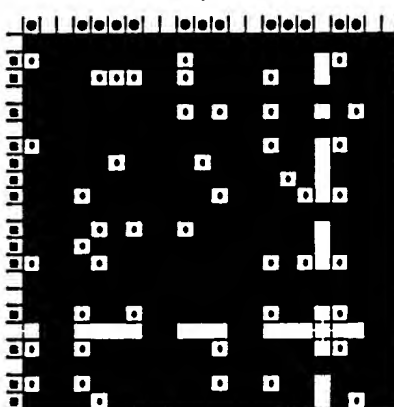


FIG.11(c)

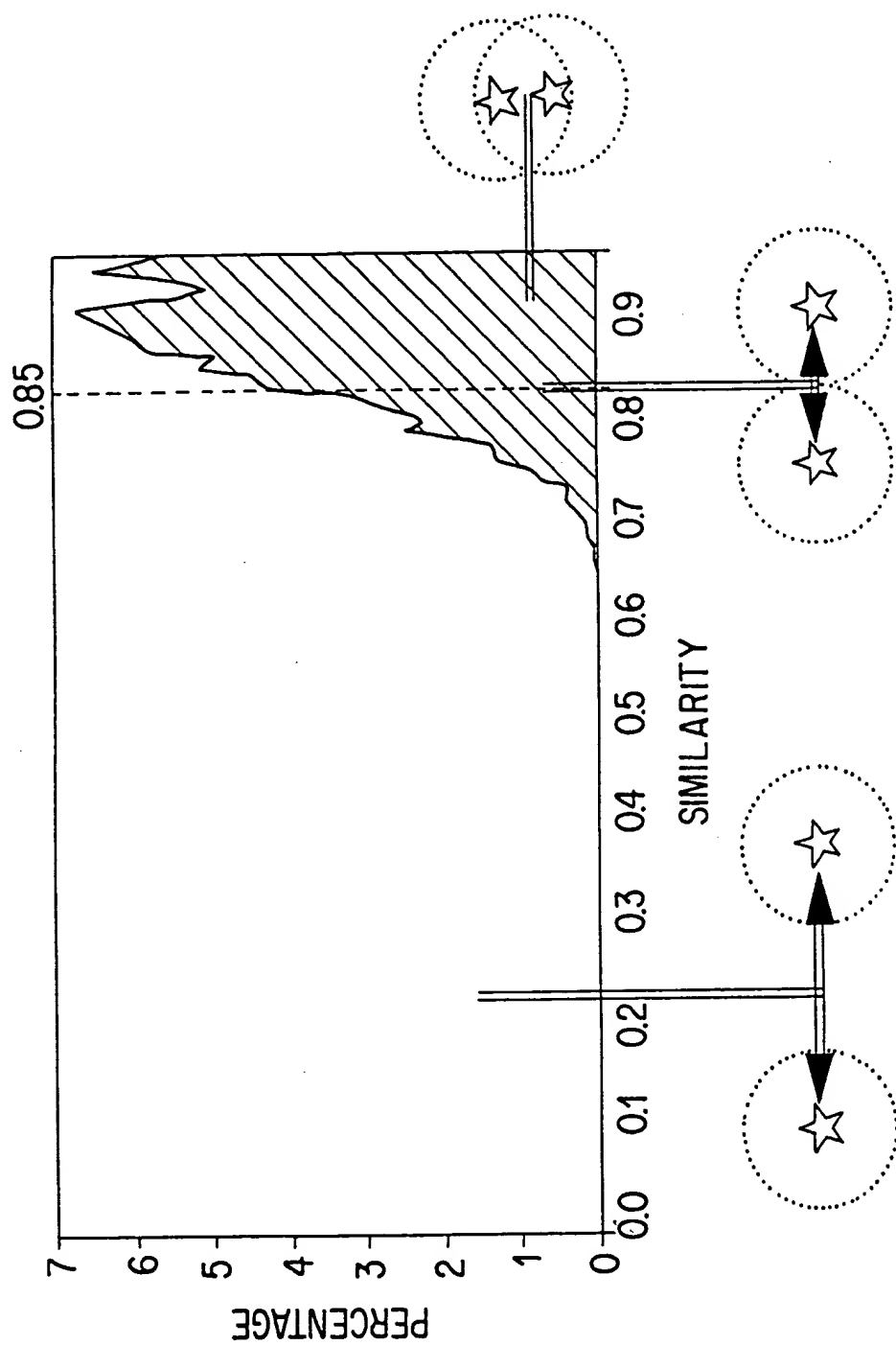


FIG.13

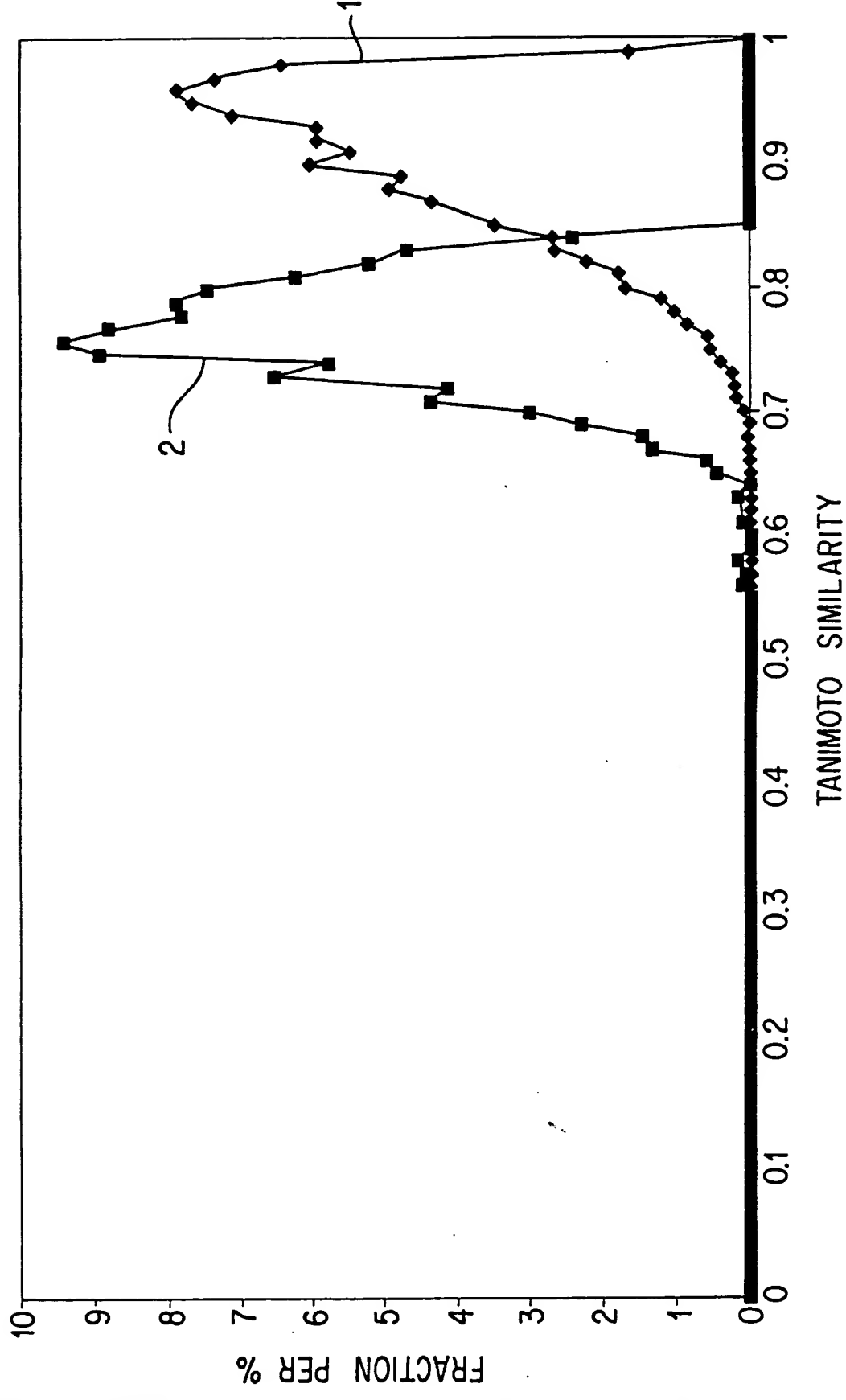


FIG. 14

